AMENDMENTS TO THE CLAIMS

What is claimed is:

What is claimed is:

1. (Currently Amended) A compound of the formula

$$R-NH-Q$$
 (I)

wherein

Q is a radical in which R₃ is hydrogen, halogen, alkyl, cycloalkyl, aryl, alkoxy, cycloalkexy, arylexy, alkylthio, cycloalkylthio, arylthio, acyl, sulfonyl, alkylamino, cycloalkylamino, arylamino, acylamino, sulfonamido or alkoxycarbonyl; Y is CH or nitrogen; and

R is a radical of the formula

wherein

R₄ is C₂₋₄alkyl, C₃₋₇cycloalkyl or C₅₋₇heterocycloalkyl;

 R_5 and R_6 are independently hydrogen, halogen, cyano, R_7 , -C(O) R_7 or -S(O) $_2$ R_7 wherein R_7 is -(CR $_8$ R $_9$) $_m$ -W-R $_{10}$ in which

R₈ and R₉ are independently hydrogen or lower alkyl;

W is a bond, O, S or NR,4 in which

Ru is hydrogen or lower alkyl;

R₁₀ is hydrogen, alkyl, cycloalkyl, aryl or heterocyclyl; or R₁₀ and R₁₁, combined, are alkylene which together with the nitrogen atom to which they are attached form a 5- to 7-membered ring:

m is zero or an integer from 1 to 5;

n is zero or an integer of 1 or 2;

or an optical isomer thereof; or a pharmaceutically acceptable salt thereof; or

(ii) Q is a radical in which R₃ is hydrogen, halogen, alkyl, cycloalkyl, aryl,

alkoxy, cycloalkoxy, aryloxy, alkylthio, cycloalkylthio, arylthio, acyl, sulfonyl, alkylamino, cycloalkylamino, arylamino, acylamino, sulfonamido or alkoxycarbonyl; and

R is a radical of the formula

wherein

R₄ is C₂₋₄alkyl, C₃₋₇cycloalkyl or C₅₋₇heterocycloalkyl;

 R_5 and R_6 are independently hydrogen, halogen, cyano, R_7 , $-C(O)R_7$ or $-S(O)_2R_7$ wherein R_7 is $-(CR_8R_9)_m$ -W-R₁₀ in which

R₈ and R₉ are independently hydrogen or lower alkyl;

W is a bond, O, S or NR₁₁ in which

R₁₁ is hydrogen or bwer alkyl;

R₁₀ is hydrogen, alkyl, cycloalkyl, aryl or heterocyclyl; or R₁₀-and R₁₄, combined, are alkylene which together with the nitrogen atom to which they are attached form a 5- to 7-membered ring;

m is zero or an integer from 1 to 5;

n is zero or an integer of 1 or 2;

or an optical isomer thereof; or a pharmaceutically acceptable salt thereof; or

(iii) Q is a

radical in which R3 is hydrogen, halogen, alkyl, cycloalkyl, aryl,

alkoxy, cycloalkoxy, aryloxy, alkylthio, cycloalkylthio, arylthio, acyl, sulfonyl, alkylamino, cycloalkylamino, arylamino, acylamino, sulfonamido or alkoxycarbonyl; and

R is a radical of the formula

wherein

R4 is C24alkyl, C37cycloalkyl or C52heterocycloalkyl;

R_s-and R_s-are independently hydrogen, halogen, cyano, R_z, -C(O)R_z or -S(O)₂R_z wherein R_z is -(CR_sR_s)_m-W-R₁₀ in which

R₈-and-R₉-are independently hydrogen or lower alkyl;

W is a bond, O, S or NR, in which

R_{1.1} is hydrogen or bwor alkyl;

R₁₀ is hydrogen, alkyl, cycloalkyl, aryl or heterocyclyl; or R₁₀ and R₁₁, combined, are alkylene which together with the nitrogen atom to which they are attached form a 5- to 7-membered ring;

m is zero or an integer from 1 to 5;

n is zero or an integer of 1 or 2;

provided that: (1) R_6 -and R_6 -are not halogen when n is zero; or (2) R_6 -is not $-S(O)_2R_7$, wherein R_7 -is $-(CR_8R_9)_m$ -W-R₁₀ in which m is zero, W is a bond and R_{10} is C_{1-3} alkyl-when n is zero;

or an optical isomer thereof; or a pharmaceutically-acceptable salt-thereof; or

R is a radical of the formula

wherein

R4 is C24alkyl, C3.zcycloalkyl or C5.zheterocycloalkyl;

R₁₂-and-R₁₃-are independently hydrogen, halogen, cyano, R₁₄, -C(O)R₁₄, or -S(O)₂R₁₄ wherein

R₁₄-is-(CR₈R₉)_m-W-R₁₅-in-which

R₈ and R₉ are independently hydrogen or lower alkyl;

Wis a bond, O, S or NR11 in which

R₁₁ is hydrogen or lower alkyl;

R₁₅ is cycloalkyl, aryl or heterocyclyl; or R₁₅ and R₁₄, combined, are alkylene which together with the nitrogen atom to which they are attached form a 5- to 7-membered ring;

m is zero or an integer from 1 to 5;

n is zero or an integer of 1 or 2;

provided that: (1) R_{12} and R_{13} both are not hydrogen, halogen, cyano or combinations thereof; (2) R_{12} is not $-S(O)_2R_{14}$, wherein R_{14} is $-(CR_8R_9)_m$ -W- R_{15} in which m is zero and W is a bond when n is zero; (3) R_{12} is not $-S(O)_2R_{14}$, wherein R_{14} is $-(CR_8R_9)_m$ -W- R_{15} in which R_8 and R_9 are hydrogen, m is 1 and W is a bond when n is zero; (4) R_{12} is not R_{14} , wherein R_{14} is $-(CR_8R_9)_m$ -W- R_{15} in which m is zero and W is O when n is zero; or (5) R_{12} is not R_{147} wherein R_{14} is $-(CR_8R_9)_m$ -W- R_{15} in which m is zero and W is a bond when n is zero; or an optical isomer thereof; or a pharmaceutically acceptable salt thereof.

2-3. (Cancelled)

(Currently Amended) A compound according to Claim 1 of the formula

$$R_{6}$$
 $(CH_{2})_{n}$
 R_{5}
 R_{4}
 $(CH_{2})_{n}$
 $(CH_{2})_{$

wherein

R₃ is hydrogen, halogen, alkyl, cycloalkyl, aryl, alkoxy, cycloalkoxy, aryloxy, alkylthio, cycloalkylthio, arylthio, acyl, sulfonyl, alkylamino, cycloalkylamino, arylamino, acylamino, sulfonamido or alkoxycarbonyl;

R₄ is C₂₋₄alkyl, C₃₋₇cycloalkyl or C₅₋₇heterocycloalkyl;

 R_5 and R_6 are independently hydrogen, halogen, cyano, R_7 , $-C(O)R_7$ or $-S(O)_2R_7$ wherein R_7 is $-(CR_8R_9)_m$ -W-R₁₀ in which

R₈ and R₉ are, independently, hydrogen or lower alkyl;

W is a bond, O, S or -NR₁₄ in which R₁₄ is hydrogen or lower alkyl;

R₁₀ is hydrogen, alkyl, cycloalkyl, aryl or heterocyclyl; or R₁₀-and R₁₁, combined, are alkylene which together with the nitrogen atom to which they are attached form a 5- to 7-membered ring;

m is zero or an integer from 1 to 5;

Y is CH-or nitrogen;

n is zero or an integer of 1 or 2;

or an optical isomer thereof; or a pharmaceutically acceptable salt thereof.

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 (Original) A compound according to Claim 4, wherein R₄ is cyclopentyl;

n is zero;

or an optical isomer thereof; or a pharmaceutically acceptable salt thereof.

6. (Currently Amended) A compound according to Claim 1 of the formula

$$R_6$$
 (CH₂) N N (Ic)

wherein

R₃ is hydrogen, halogen, alkyl, cyclealkyl, aryl, alkoxy, cyclealkoxy, arylexy, alkylthie, cyclealkylthie, arylthie, acyl, sulfenyl, alkylamine, cyclealkylamine, arylamine, acylamine, sulfenamide or alkoxycarbenyl;

R₄ is C₂₋₄alkyl, C₃₋₇cycloalkyl or C₅₋₇heterocycloalkyl;

 R_5 and R_6 are independently hydrogen, halogen, cyano, R_7 , -C(O) R_7 or -S(O) $_2R_7$ wherein R_7 is -(CR $_8R_9$) $_m$ -W-R $_{10}$ in which

R₈ and R₉ are, independently, hydrogen or lower alkyl;

W is a bond, O, S or NR, in which

R₁₄ is hydrogen or lower-alkyl;

R₁₀ is hydrogen, alkyl, cycloalkyl, aryl or heterocyclyl; or R₁₀-and R₁₄, combined, are alkylene which together with the nitrogen atom to which they are attached form a 5-to-7-membered ring;

m is zero or an integer from 1 to 5;

n is zero or an integer of 1 or 2;

or an optical isomer thereof; or a pharmaceutically acceptable salt thereof.

7. (Original) A compound according to Claim 6, wherein

R₄ is cyclopentyl;

n is zero;

or an optical isomer thereof; or a pharmaceutically acceptable salt thereof.

8-11. (Cancelled)

- 12. (Withdrawn) A method for the activation of glucokinase activity in mammals which method comprises administering to a mammal in need thereof a therapeutically effective amount of a compound of claim 1.
- 13. (Withdrawn) A method for the prevention and/or treatment of conditions associated with glucokinase activity in mammals which method comprises administering to a mammal in need thereof a therapeutically effective amount of a compound of claim 1.
- 14. (Withdrawn) The method according to claim 13, which method comprises administering said compound in combination with a therapeutically effective amount of insulin, insulin derived mimetic; insulin secretagogue; insulinotropic sulfonylurea receptor ligand; PPAR ligand; insulin sensitizer; biguanide; alpha-glucose inhibitors; GLP-1, GLP-1 analog or mimetic; DPPIV inhibitor; HMG-CoA reductase inhibitor; squaline synthase inhibitor; FXR or LXR ligand; cholestyramine; fibrates; nicotinic acid; or aspirin.
- 15. (Withdrawn) A method for the treatment of impaired glucose tolerance, Type 2 diabetes and obesity which method comprises administering to a mammal in need thereof a therapeutically effective amount of a compound of claim 1.
- 16. (Original) A pharmaceutical composition comprising a therapeutically effective amount of a compound of claims 1 in combination with one or more pharmaceutically acceptable carriers.
- 17. (Original) A pharmaceutical composition comprising a therapeutically effective amount of a compound of claim 1 in combination with a pharmaceutically effective amount of insulin, insulin derived mimetic; insulin secretagogue; insulinotropic sulfonylurea receptor ligand; PPAR ligand; insulin sensitizer; biguanide; alpha-glucose inhibitors; GLP-1, GLP-1 analog or mimetic; DPPIV inhibitor; HMG-CoA reductase inhibitor; squaline synthase inhibitor; FXR or LXR ligand; cholestyramine; fibrates; nicotinic acid; or aspirin.
- 18. (Previously Presented) A paharmaceutical composition according to claim 16 for the treatment of impaired glucose tolerance, Type 2 diabetes and obesity.
- 19 24. (Cancelled)

25. (New) A compound according to Claim 6, wherein the compound is: